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PTO/SB/21 (09-04) Approved for use through 07/31/2006. OMB 0651-0031
U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE he Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number. Patent Number 6,770,486 TRANSMITTAL Issue Date 08/03/2004 First Named Inventor **FORM** Richard Griffey OPTIMIZATION OF LIGAND AFFINITY FOR RNA TARGETS Title USING MASS SPECTROMETRY (to be used for all correspondence after initial filing) Attorney Docket Number MSIBIS-0003US Total Number of Pages in This Submission **ENCLOSURES** (Check all that apply) After Allowance Communication to TC Fee Transmittal Form Drawing(s) Appeal Communication to Board Licensing-related Papers Fee Attached of Appeals and Interferences Appeal Communication to TC Petition (Appeal Notice, Brief, Reply Brief) Amendment/Reply Petition to Convert to a Proprietary Information After Final **Provisional Application** Power of Attorney, Revocation Status Letter Affidavits/declaration(s) Change of Correspondence Address Other Enclosure(s) (please Identify **Terminal Disclaimer Extension of Time Request** below): Certificate of Correction Request for Refund Express Abandonment Request CD, Number of CD(s) Information Disclosure Statement Certificate Landscape Table on CD Certified Copy of Priority Remarks Document(s) FEB 1 6 2006 Reply to Missing Parts/ Incomplete Application of Correction Reply to Missing Parts under 37 CFR 1.52 or 1.53 SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT Firm Name Isis Pharmaceuticals, Inc. Signature Printed name Herb Boswell Date Reg. No. 27,311 60 CERTIFICATE OF TRANSMISSION/MAILING I hereby certify that this correspondence is being facsimile transmitted to the USPTO or deposited with the United States Postal Service with sufficient postage as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450 on the date shown below: Signature Date Kemlyn Evans Typed or printed name

This collection of information is required by 37 CFR 1.5. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.11 and1.14. This collection is estimated to 2 hours to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

UNITED STATES PATENT AND TRADEMARK OFFICE CERTIFICATE OF CORRECTION

PATENT NO. : 6,770,486

DATED

: 08/03/2004

INVENTOR(S) : Richard Griffey et al.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

1)Title Page:

Item [56], References Cited, OTHER PUBLICATIONS, "Griffey" reference, please delete "mismtches" and insert therefor -- mismatches --;

2) Title Page:

Item [74], Attorney, Agent, or Firm, please delete "Rosch" and insert therefor -- Roach --;

3) Column 31, Claim 1, line 46, please delete "noncovalent" and insert therefor -- non-covalent --.

MAILING ADDRESS OF SENDER:

PATENT NO. 6,770,486

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